

In the Specification:

Please insert the following paragraph at page 1, line 2:

**--RELATED APPLICATIONS**

This is a continuation application claiming priority under 35 USC §120 to US serial number 09/665,350 filed 9/18/2000, which is a continuation of, and claims priority under 35 USC §120 to, PCT Application number PCT/US00/04414 filed 2/22/2000, and where US serial number 09/665,350 is also a continuation-in-part of, and claims priority under 35 USC §120 to, PCT Application number PCT/US00/23328 filed 8/24/2000, which is a continuation-in-part of, and claims priority under 35 USC §120 to, PCT Application number PCT/US00/20710 filed 7/28/2000, which claims priority under 35 USC §119 to US provisional application number 60/146,222 filed 7/28/1999, and where PCT Application number PCT/US00/20710 is also a continuation-in-part of, and claims priority under 35 USC §120 to, PCT Application number PCT/US00/15264 filed 6/2/2000, which is a continuation-in-part of, and claims priority under 35 USC §120 to, PCT Application number PCT/US00/14042 filed 5/22/2000, which is a continuation-in-part of, and claims priority under 35 USC §120 to, PCT Application number PCT/US00/08439 filed 3/30/2000, which is a continuation-in-part of, and claims priority under 35 USC §120 to, PCT Application number PCT/US00/07377 filed 3/20/2000, which is a continuation-in-part of, and claims priority under 35 USC §120 to, PCT Application number PCT/US00/05841 filed 3/2/2000, which is a continuation-in-part of, and claims priority under 35 USC §120 to, PCT Application number PCT/US00/05004 filed 2/24/2000, which is a continuation-in-part of, and claims priority under 35 USC §120 to, PCT Application number PCT/US00/04414 filed 2/22/2000, which is a continuation-in-part of, and claims priority under 35 USC §120 to, PCT Application number PCT/US00/03565 filed 2/11/2000, which is a continuation-in-part of, and claims priority under 35 USC §120 to, PCT Application number PCT/US00/00219 filed 1/5/2000, which claims priority under 35 USC §119 to US provisional application number 60/145,698 filed 7/26/1999, and where PCT Application number PCT/US00/00219 is also a continuation-in-part of, and claims priority under 35 USC §120 to, both PCT Application numbers: PCT/US99/30999 filed 12/20/1999 and PCT/US99/30911 filed 12/20/1999, which claims priority under 35 USC §119 to US provisional application number 60/143,048 filed 7/7/1999, and where PCT/US99/30999 and PCT/US99/30911 are both continuations-in-part to, and claim priority under 35 USC §120 to, PCT Application number

PCT/US99/30095 filed 12/16/1999, which claims priority under 35 USC §119 to US provisional application number 60/113,296 filed 12/22/1998, and where PCT Application number PCT/US99/30095 is also a continuation-in-part of, and claims priority under 35 USC §120 to, both PCT Application numbers: PCT/US99/28565 and PCT/US99/28564 both filed 12/2/1999, which are also continuations-in-part of, and claim priority under 35 USC §120 to, PCT Application number PCT/US99/28301 filed 12/1/1999, which is a continuation-in-part of, and claims priority under 35 USC §120 to, PCT Application number PCT/US99/28313 filed 11/30/1999, which is a continuation-in-part of, and claims priority under 35 USC §120 to, PCT Application number PCT/US99/28214 filed 11/29/1999, which is a continuation-in-part of, and claims priority under 35 USC §120 to, PCT Application number PCT/US99/23089 filed 10/5/1999, which claims priority under 35 USC §119 to US provisional application number 60/104080 filed 10/13/1998, and where PCT Application number PCT/US99/23089 is also a continuation-in-part of, and claims priority under 35 USC §120 to, both PCT Application numbers: PCT/US99/21547 and PCT/US99/21090 both filed 9/15/1999, which are also continuations-in-part of, and claim priority under 35 USC §120 to, PCT Application number PCT/US99/20944 filed 9/13/1999, which is a continuation-in-part of, and claims priority under 35 USC §120 to, PCT Application number PCT/US99/20594 filed 9/8/1999, which is a continuation-in-part of, and claims priority under 35 USC §120 to, PCT Application number PCT/US98/25108 filed 12/1/1998, which is a continuation-in-part of, and claims priority under 35 USC §120 to, PCT Application number PCT/US98/19437 filed 9/17/1998, which is a continuation-in-part of, and claims priority under 35 USC §120 to, PCT Application number PCT/US98/19330 filed 9/16/1998, which is a continuation-in-part of, and claims priority under 35 USC §120 to, PCT Application number PCT/US98/19177 filed 9/14/1998, which is a continuation-in-part of, and claims priority under 35 USC §120 to, PCT Application number PCT/US98/18824 filed 9/10/1998, which claims priority under 35 USC §119 to US provisional application numbers: 60/059,115 filed 9/17/1997, 60/059,184 filed 9/17/1997, 60/059,122 filed 9/17/1997, 60/059,117 filed 9/17/1997, 60/059,113, filed 9/17/1997, 60/059,121 filed 9/17/1997, 60/059,119 filed 9/17/1997, 60/059,263 filed 9/18/1997, 60/059,266 filed 9/18/1997, 60/062,125 filed 10/15/1997, 60/062,287 filed 10/17/1997, 60/062,285 filed 10/17/1997, 60/063,486 filed 10/21/1997, 60/062,816 filed 10/24/1997, 60/062,814 filed 10/24/1997, 60/063,127 filed 10/24/1997, 60/063,120 filed 10/24/1997, 60/063,121 filed 10/24/1997, 60/063,045 filed

10/24/1997, 60/063,128 filed 10/24/1997, 60/063,329 filed 10/27/1997, 60/063,327 filed 10/27/1997, 60/063,549 filed 10/28/1997, 60/063,541 filed 10/28/1997, 60/063,550 filed 10/28/1997, 60/063,542 filed 10/28/1997, 60/063,544 filed 10/28/1997, 60/063,564 filed 10/28/1997, 60/063,734 filed 10/29/1997, 60/063,738 filed 10/29/1997, 60/063,704 filed 10/29/1997, 60/063,435 filed 10/29/1997, 60/064,215 filed 10/29/1997, 60/063,735 filed 10/29/1997, 60/063,732 filed 10/29/1997, 60/064,103 filed 10/31/1997, 60/063,870 filed 10/31/1997, 60/064,248 filed 11/3/1997, 60/064,809 filed 11/7/1997, 60/065,186 filed 11/12/1997, 60/065,846 filed 11/17/1997, 60/065,693 filed 11/18/1997, 60/066,120 filed 11/21/1997, 60/066,364 filed 11/21/1997, 60/066,772 filed 11/24/1997, 60/066,466 filed 11/24/1997, 60/066,770 filed 11/24/1997, 60/066,511 filed 11/24/1997, 60/066,453 filed 11/24/1997, 60/066,840 filed 11/25/1997, 60/069,425 filed 12/12/1997, 60/088,026 filed 6/4/1998, 60/109,304 filed 11/20/1998, the entire disclosures of which are hereby incorporated by reference.

The paragraph, beginning at page 69, line 6, has been amended as follows:

--Percent amino acid sequence identity may also be determined using the sequence comparison program NCBI-BLAST2 (Altschul et al., Nucleic Acids Res. 25:3389-3402 (1997)).  
The NCBI-BLAST2 sequence comparison program may be downloaded from <http://www.ncbi.nlm.nih.gov>. NCBI-BLAST2 uses several search parameters, wherein all of those search parameters are set to default values including, for example, unmask = yes, strand = all, expected occurrences = 10, minimum low complexity length = 15/5, multi-pass e-value = 0.01, constant for multi-pass = 25, dropoff for final gapped alignment = 25 and scoring matrix = BLOSUM62.--

The paragraph, beginning at page 71, line 26, has been amended as follows:

--Percent nucleic acid sequence identity may also be determined using the sequence comparison program NCBI-BLAST2 (Altschul et al., Nucleic Acids Res. 25:3389-3402 (1997)).  
The NCBI-BLAST2 sequence comparison program may be downloaded from <http://www.ncbi.nlm.nih.gov>. NCBI-BLAST2 uses several search parameters, wherein all of those search parameters are set to default values including, for example, unmask = yes, strand = all, expected occurrences = 10, minimum low complexity length = 15/5, multi-pass e-value =

0.01, constant for multi-pass = 25, dropoff for final gapped alignment = 25 and scoring matrix = BLOSUM62.--

The paragraph, beginning at page 147, line 20 has been amended as follows:

--Commercially available reagents referred to in the examples were used according to manufacturer's instructions unless otherwise indicated. The source of those cells identified in the following examples, and throughout the specification, by ATCC accession numbers is the American Type Culture Collection, [Rockville, Maryland]Manassas, VA--

The paragraph beginning at page 147, line 27, has been amended as follows:

--The extracellular domain (ECD) sequences (including the secretion signal sequence, if any) from about 950 known secreted proteins from the Swiss-Prot public database were used to search EST databases. The EST databases included public databases (e.g., Dayhoff, GenBank), and proprietary databases (e.g. LIFESEQ™, Incyte Pharmaceuticals, Palo Alto, CA). The search was performed using the computer program BLAST or BLAST2 (Altschul, and Gish, Methods in Enzymology 266: 460-80 (1996); <http://blast.wustl.edu/blast/README.html>) as a comparison of the ECD protein sequences to a 6 frame translation of the EST sequences. Those comparisons with a Blast score of 70 (or in some cases 90) or greater that did not encode known proteins were clustered and assembled into consensus DNA sequences with the program "phrap" (Phil Green, University of Washington, Seattle, Washington).

The paragraph, beginning at page 154, line 14 has been amended as follows:

--The EST sequence accession number AF007268, a murine fibroblast growth factor (FGF-15) was used to search various public EST databases (e.g., GenBank, Dayhoff, etc.) The search was performed using the computer program BLAST or BLAST2 Altsehul et al., Methods in Enzymology, 266:460-480 (1996); <http://blast.wustl.edu/blast/README.html> as a comparison of the ECD protein sequences to a 6 frame translation of the EST sequences. The search resulted in a hit with GenBank EST AA220994, which has been identified as stratagene Strategene NT2 neuronal precursor 937230.--

The paragraph beginning at page 167, line 30, has been amended as follows:

--The extracellular domain (ECD) sequences (including the secretion signal, if any) of from about 950 known secreted proteins from the Swiss-Prot public protein database were used to search expressed sequence tag (EST) databases. The EST databases included public EST databases (e.g., GenBank) and a proprietary EST DNA database (LIFESEQ™, Incyte Pharmaceuticals, Palo Alto, CA). The search was performed using the computer program BLAST or BLAST2 (Altshul et al., Methods in Enzymology 266:460-480 (1996)) as a comparison of the ECD protein sequences to a 6 frame translation of the EST sequence. Those comparisons resulting in a BLAST score of 70 (or in some cases 90) or greater that did not encode known proteins were clustered and assembled into consensus DNA sequences with the program "phrap" (Phil Green, University of Washington, Seattle, Washington; [http://bozeman.mbt.washington.edu/phrap.docs/phrap.html](http://bozeman.mbt.washington.edu/phrap/docs/phrap.html))--.

The paragraph beginning at page 178, line 26, has been amended as follows:

--The extracellular domain (ECD) sequences (including the secretion signal, if any) of from about 950 known secreted proteins from the Swiss-Prot public protein database were used to search expressed sequence tag (EST) databases. The EST databases included public EST databases (e.g., GenBank) and a proprietary EST DNA database (LIFESEQ™, Incyte Pharmaceuticals, Palo Alto, CA). The search was performed using the computer program BLAST or BLAST2 (Altshul et al., Methods in Enzymology 266:460-480 (1996)) as a comparison of the ECD protein sequences to a 6 frame translation of the EST sequence. Those comparisons resulting in a BLAST score of 70 (or in some cases 90) or greater that did not encode known proteins were clustered and assembled into consensus DNA sequences with the program "phrap" (Phil Green, University of Washington, Seattle, Washington; [http://bozeman.mbt.washington.edu/phrap.docs/phrap.html](http://bozeman.mbt.washington.edu/phrap/docs/phrap.html)).--

The paragraph, beginning at page 250, line 2, has been amended as follows:

--The following materials have been deposited with the American Type Culture Collection, [12301 Parklawn Drive, Rockville, MD,] 10801 University Boulevard, Manassas, VA USA (ATCC):

| <u>Material</u> | <u>ATCC Dep. No.</u> | <u>Deposit Date</u> |
|-----------------|----------------------|---------------------|
| DNA32292-1131   | ATCC 209258          | September 16, 1997  |
| DNA33094-1131   | ATCC 209256          | September 16, 1997  |
| DNA33223-1136   | ATCC 209264          | September 16, 1997  |
| DNA34435-1140   | ATCC 209250          | September 16, 1997  |
| DNA27864-1155   | ATCC 209375          | October 16, 1997    |
| DNA36350-1158   | ATCC 209378          | October 16, 1997    |
| DNA32290-1164   | ATCC 209384          | October 16, 1997    |
| DNA35639-1172   | ATCC 209396          | October 17, 1997    |
| DNA33092-1202   | ATCC 209420          | October 28, 1997    |
| DNA49435-1219   | ATCC 209480          | November 21, 1997   |
| DNA35638-1141   | ATCC 209265          | September 16, 1997  |
| DNA32298-1132   | ATCC 209257          | September 16, 1997  |
| DNA33089-1132   | ATCC 209262          | September 16, 1997  |
| DNA33786-1132   | ATCC 209253          | September 16, 1997  |
| DNA35918-1174   | ATCC 209402          | October 17, 1997    |
| DNA37150-1178   | ATCC 209401          | October 17, 1997    |
| DNA38260-1180   | ATCC 209397          | October 17, 1997    |
| DNA39969-1185   | ATCC 209400          | October 17, 1997    |
| DNA32286-1191   | ATCC 209385          | October 16, 1997    |
| DNA33461-1199   | ATCC 209367          | October 15, 1997    |
| DNA40628-1216   | ATCC 209432          | November 7, 1997    |
| DNA33221-1133   | ATCC 209263          | September 16, 1997  |
| DNA33107-1135   | ATCC 209251          | September 16, 1997  |
| DNA35557-1137   | ATCC 209255          | September 16, 1997  |
| DNA34434-1139   | ATCC 209252          | September 16, 1997  |
| DNA33100-1159   | ATCC 209373          | October 16, 1997    |
| DNA35600-1162   | ATCC 209370          | October 16, 1997    |
| DNA34436-1238   | ATCC 209523          | December 10, 1997   |
| DNA33206-1165   | ATCC 209372          | October 16, 1997    |
| DNA35558-1167   | ATCC 209374          | October 16, 1997    |
| DNA35599-1168   | ATCC 209373          | October 16, 1997    |
| DNA36992-1168   | ATCC 209382          | October 16, 1997    |
| DNA34407-1169   | ATCC 209383          | October 16, 1997    |
| DNA35841-1173   | ATCC 209403          | October 17, 1997    |
| DNA33470-1175   | ATCC 209398          | October 17, 1997    |
| DNA34431-1177   | ATCC 209399          | October 17, 1997    |
| DNA39510-1181   | ATCC 209392          | October 17, 1997    |
| DNA39423-1182   | ATCC 209387          | October 17, 1997    |
| DNA40620-1183   | ATCC 209388          | October 17, 1997    |
| DNA40604-1187   | ATCC 209394          | October 17, 1997    |
| DNA38268-1188   | ATCC 209421          | October 28, 1997    |
| DNA37151-1193   | ATCC 209393          | October 17, 1997    |
| DNA35673-1201   | ATCC 209418          | October 28, 1997    |
| DNA40370-1217   | ATCC 209485          | November 21, 1997   |
| DNA42551-1217   | ATCC 209483          | November 21, 1997   |

|               |             |                     |
|---------------|-------------|---------------------|
| DNA39520-1217 | ATCC 209482 | November 21, 1997   |
| DNA41225-1217 | ATCC 209491 | November 21, 1997   |
| DNA43318-1217 | ATCC 209481 | November 21, 1997   |
| DNA40587-1231 | ATCC 209438 | November 7, 1997    |
| DNA41338-1234 | ATCC 209927 | June 2, 1998        |
| DNA40981-1234 | ATCC 209439 | November 7, 1997    |
| DNA37140-1234 | ATCC 209489 | November 21, 1997   |
| DNA40982-1235 | ATCC 209433 | November 7, 1997    |
| DNA41379-1236 | ATCC 209488 | November 21, 1997   |
| DNA44167-1243 | ATCC 209434 | November 7, 1997    |
| DNA39427-1179 | ATCC 209395 | October 17, 1997    |
| DNA40603-1232 | ATCC 209486 | November 21, 1997   |
| DNA43466-1225 | ATCC 209490 | November 21, 1997   |
| DNA43046-1225 | ATCC 209484 | November 21, 1997   |
| DNA35668-1171 | ATCC 209371 | October 16, 1997    |
| DNA77624-2515 | ATCC 203553 | December 22, 1998-- |

Please replace the paragraph beginning at page 251, line 10, with the following rewritten paragraph:

--These deposit were made under the provisions of the Budapest Treaty on the International Recognition of the Deposit of Microorganisms for the Purpose of Patent Procedure and the Regulations there under (Budapest Treaty). This assures maintenance of a viable culture of the deposit for 30 years from the date of deposit. The deposits will be made available by ATCC under the terms of the Budapest Treaty, and subject to an agreement between Genentech, Inc. and ATCC, which assures that all restrictions imposed by the depositor on the availability to the public of the deposited material will be irrevocably removed upon the granting of the pertinent U.S. patent, assures permanent and unrestricted availability of the progeny of the culture of the deposit to the public upon issuance of the pertinent U.S. patent or upon laying open to the public of any U.S. or foreign patent application, whichever comes first, and assures availability of the progeny to one determined by the U.S. Commissioner of Patents and Trademarks to be entitled thereto according to 35 USC § 122 and the Commissioner's rules pursuant thereto (including 37 CFR § 1.14 with particular reference to 886 OG 638).--